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The demark Office is displays a valid OMB control Please type a plus sign (+) inside this box -> + Approved for use through +0x1/2005 10 ME das 100 ME C 3. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of months in the description unless it displays a valid OMB control number. 09/888,309 Applicati n Number TRANSMITTAL June 21, 2001 Filing Date **FORM** First Named Inventor Melissa K. Carpenter **Group Art Unit** (to be used for all correspondence after initial filing) 1632 Ton, Thaian N. **Examiner Name** 090/002 Total Number of Pages in This Submission Attorney Docket Number **ENCLOSURES** (check all that apply) After Allowance Communication Assignment Papers Fee Transmittal Form (for an Application) to Group Appeal Communication to Board Fee Attached Drawing(s) of Appeals and Interferences Appeal Communication to Group Licensing-related Papers Amendment / Reply (Appeal Notice, Brief, Reply Brief) Petition After Final Proprietary Information Petition to Convert to a Affidavits/declaration(s) Provisional Application Status Letter Power of Attorney, Revocation Change of Correspondence Address Other Enclosure(s) (please Extension of Time Request identify below): Terminal Disclaimer PTO Form 1449; References (56) **Express Abandonment Request** Request for Refund Information Disclosure Statement CD, Number of CD(s). Certified Copy of Priority Document(s) Remarks Response to Missing Parts/ Incomplete Application Response to Missing Parts under 37 CFR 1.52 or 1.53 SIGNATURE OF APPLICANT, ATTORNEY, OR AGENT Firm J. Michael Schiff, Registration No. 40,253 Individual name Signature Date Noumber 30, 2001 CERTIFICATE OF MAILING I hereby certify that this correspondence is being deposited with the United States Postal Service with sufficient postage as first class mail in an envelope addressed to: Commissioner for Patents, Washington, DC 20231 on this date: Typed or printed name

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I hereby certify that this correspondence is being delivered by hand to the U.S. Patent Office in accordance with 37 CFR § 1.6(b), addressed to the Assistant Commissioner for Patents, Washington, D.C. 20231, on the date indicated.

Name

Date

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In the application of: Carpenter et al.

Serial No.: 09/888,309

Filing Date: June 21, 2001

For: DIRECT DIFFERENTIATION OF HUMAN

PLURIPOTENT STEM CELLS AND

CHARACTERIZATION OF DIFFERENTIATED

CELLS

Art Unit: 1632

Examiner: Ton, Thaian N.

TECH CENTER 1600/290

INFORMATION DISCLOSURE STATEMENT

Assistant Commissioner for Patents Washington, D.C. 20231

Dear Sir:

The information listed in the accompanying form PTO-1449 and provided herewith may be material to examination of this application and is submitted in compliance with the duty of disclosure under 37 CFR § 1.56. The Examiner is requested to make this information of record in the application.

Docket: 090/002

This Information Disclosure Statement is not to be construed as a representation that a full search for relevant information has been made, that all relevant information has been found, or that the information provided with this Statement is considered to be material to patentability of the claimed invention as defined under 37 CFR § 1.56(b).

It is believed that no fee is required for submission of this Statement, which is filed before the first Office Action on the merits of the application. Nevertheless, should a fee be required for consideration of this Statement and the listed information, the Assistant Commissioner is authorized to charge such fee to Deposit Account No. 07-1139, referencing the attorney Docket Number indicated above.

Respectfully submitted,

J. Midhael Schiff

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DATE: November **28**, 2001

F rm 1449 (modified)

Inf rmati n Disci sure Stat m nt By Applicant

(Use Several Sheets if Necessary)

Docket: 090/002

U. N. 09/888,309

Title: Direct Differentiation of Human Pluripotent Stem Cells and Characterization of Differentiated Cells
Inventors: Melissa K. Carpenter, Walter D. Funk, R. Scott Theis

Solution 21, 2001

Group: 1632

Examiner Initial	Ref.	Patent No.	Filing Date	Issue Date	Class/ Subclass	Inventors:	Title:
	Α	5,766,948	Nov 3/93	Jun 16/98	435/368	Gage, F.H., et al.	Method for Production of Neuroblasts
	В	5,773,255	Jun 5/95	Jun 30/98	435/70.3	Laurance, M.E., et al.	Glucose Responsive Insulin Secreting β-Cell Lines and Method For Producing Same
	С	5,789,246	Nov 18/96	Aug 4/98	435/325	Reid, L.M., et al.	Compositions Comprising Hepatocyte Precursors
	D	5,849,553	Jun 7/95	Dec 15/98	435/172.3	Anderson, D.J., et al.	Mammalian Multipotent Neural Stem Cells
	E	5,851,832	Jun 7/95	Dec 22/98	435/368	Weiss, S., et al.	In Vitro Growth and Proliferation of Multipotent neural Stem Cells and Their Progeny
	F	5,928,947	Jun 7/95	Jul 27/99	435/455	Anderson, D.J., et al.	Mammalian Multipotent Neural Stem Cells
	G	5,968,829	Sep 5/97	Oct 19/99	435/467	Carpenter, M.	Human CNS Neural Stem Cells
	н	5,981,165	Jun 7/95	Nov 9/99	435/4	Weiss S., et al.	In Vitro Induction of Dopaminergic Cells
	ı	6,040,180	May 7/97	Mar 21/00	435/377	Johe, K.	In vitro Generation of Differentiated Neurons From Cultures of mammalian Multipotent CNS Stem Cells
	J	6,090,622	Mar 31/97	Jul 18/00	435/366	Gearheart, J.D., et al.	Human Embryonic Pluripotent Germ Cells
	К	6,200,806	Jun 26/98	Mar 13/01	435/366	Thomson, J.A.	Primate Embronic Stem Cells

Foreign Patent or Published Foreign Patent Application

Examiner	1 5	Document	Publ.	Juris-	Title:	Translation	
Initial	Ref.	No.	Date	diction	riue.	Yes	No
	L	WO 99/04775	Feb 4/99	PCT	Method of Treating Dopaminergic and Gaba-Nergic Disorders		
	М	WO 99/20741	Apr 29/99	PCT	Methods and Materials for the Growth of Primate- Derived Primordial Stem Cells		
	N	WO 99/43785	Sep 2/99	PCT	Derivation of Cells and Tissues from Embryonic Pre- Stem Cells for Transplantation Therapies		
	0	WO 99/53021	Oct 21/99	PCT	Cell Differentiation/Proliferation and Maintnance and Uses Thereof		
	Р	WO 00/17323	Mar 30/00	PCT	Stable Neural Stem Cell Lines		

Examiner	Date Considered

Form 1449 (modified)

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Inventors: Melissa K. Carpenter, Walter D. Funk, R. Scott Theis

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Other Documents

Examiner Initial	Ref.	Author, Title, Date, Source
	Q	Andrews, et al., Retinoic Acid Induces Neuronal Differentiation of a Cloned Haman Embryonal Carcinoma Cell Line in Vitro, Dev. Biol. 103:285 (1984)
	R	Bain, et al., Embryonic Stem Cells Express Neuronal Properties In Vitro, Dev. Biol. 168:342 (1995)
	s	Bain, et al., Expression of Retinoid X Receptors in P19 Embryonal Carcinoma Cells and Embryonic Stem Cells, Biochem. Biophys. Res. Commun. 200:1252 (1994)
	Т	Bain, et al., Retinoic Acid Promotes Neural and Represses Mesodermal Gene Expression in Mouse Embryonic Stem Cells in Culture, Chem. and Biophys. Res. Comm. 223:691 (1996)
	U	Bieseckert, et al., Interleukin-6 is a Component of Hman Umbilical Cord Serum and Stimulates Hematopoiesis in Embryonic Stem Cells in Vitro, Exp. Hematol. 21:744 (1993)
	V	Bouwmeester, et al., Vertebrate Head Induction By Anterior Primitive Endoderm, BioEssays 19:855 (1997)
	w	Brustle, et al., In Vitro-Generated Neural Precursors Participate in Mammalian Brain Development, Proc. Natl. Acad. Sci. USA 94:14809 (1997)
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	Υ	Burkert, et al., Early Fetal hematopoietic Development From In Vitro Differentiated Embryonic Stem Cells, New Biol. 3:698 (1991)
	Z	Davidson, et al., Cell Fate and Lineage Specification in the Gastrulating Mouse Embryo, Children's Medical Res. Institute :491 (1999)
	AA	Deacon, et al., Blastula-Stage Stem Cells Can Differentiate into Dopaminergic and Serotonergic Neurons after Transplantation, Exp. Neurol. 149:28 (1998)
	АВ	Dinsmore, et al., Embryonic Stem Cells Differentiated In Vitro as a Novel Source of Cells for Transplantation, Cell Transplant 5:131 (1996)
	AC	Fisher, et al., Factors Influencing the Differentiation of Embryonal Carcinoma and Embryo-Derived Stem Cells, Exp. Cell Research 182:403 (1989)
	AD	Fraichard, et al., In Vitro Differentiationof Embryonic Stem Cells into Glial Cells and Functional Neurons, J. Cell Science 108:3181 (1995)
	AE	Gendron, et al., Induction of Embryonic Vasculogenesis by bFGF and LIF In Vitro and In Vivo, Dev. Biol 177:332 (1996)
	AF	Itskovitz-Eldor, et al., Differentiation of Human Embryonic Stem Cells into Embryoid Bodies Comprising the Three Embryonic Germ Layers, Mol. Med. 6:88 (2000)
	AG	Kalyani, et al., Cell Lineage in the Developing Neural Tube, Biochem. Cell. Biol. 76:1051 (1998)
	АН	Keller, In Vitro Differentiation of Embryonic Stem Cells, Cell Biology 7:862 (1995)
	AI	Levinson-Dushnik, et al., Involvement of Hepatocyte Nuclear Factor 3 in Endoderm Differentiation of Embryonic Stem Cells, Mol. Cell. Biol. 17:3817 (1997)
	AJ	Mujtaba, et al., Lineage-Restricted Neural Precursors Can Be Isolated from Both the Mouse neural Tube and Cultured ES Cells, Dev. Biol. 214:113 (1999)
	AK	Mummery, et al., Characteristics of Embryonic Stem Cell Differentiation: A Comparison With Two Embryonal Carcinoma Cell Lines, Cell Diff. Dev. 30:195 (1990)
	AL	Odorico, et al., Multilineage Differentiation from Human Embryonic Stem Cell Lines, Stem Cells 19:193 (2001)
	AM	Okabe, et al., Development of Neuronal Precursor Cells and Functional Postmitotic Neurons from Embryonic Stem Cells In Vitro, Mech. Dev. 59:89 (1996)

Examiner	Date Considered

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Examiner Initial	Ref.	Author, Title, Date, Source
	AN	O'Shea, Embryonic Stem Cell Models of Development, Anat. Rec. (New Anat.) 257:32 (1999)
	AO	Pedersen, Studies of In Vitro Differentiation with Embryonic Stem Cells, Reprod. Fertil. Dev. 6:542 (1994)
	AP	Rao, Multipotent and Restricted Precursors in the Central Nervous System, New Anat. 257:1 (1999)
	AQ	Rathjen, et al., Formation of a Primitive Ectoderm Like Cell Population, EPL Cells, From ES Cells in Response to Biologically Derived Factors, J. Cell. Sci. 112:601 (1999)
_	AR	Rathjen, et al., Properties and Uses of Embryonic Stem Cells: Prospects for Application to Human Biology and Gene Therapy, Reprod. Fertil. Dev. 10:31 (1998)
	AS	Reubinof, et al., Embryonic Stem Cell Lines From Human Blastocysts: Somatic Differentiation In Vitro, Nature Biol. 18:399 (2000)
	AT	Robertson, Derivation and Maintenance of Embryonic Stem Cell Cultures, Meth. Mol. Biol. 75:173 (1997)
	AU	Schuldiner, et al., Effects of Eight Growth Factors on the Differentiation of Cells Derived from Human Embryonic Stem Cells, PNAS 97:11307 (2000)
	AV	Strubing, et al., Differentiationof Pluripotent Embryonic Stem Cells into the Neuronal Lineage in Vitro Gives Rise to Mature Inhibitory and Excitatory Neurons, Mechanisms of Dev. 53"275 (1995)
	AW	Seaberg, et al., Neural Determination Genes Revealed By Expression Trapping in Embryonic Stem Cells, Soc. Neurosci. (29 th Annual Meeting) 25:527 (1999)
	AX	Shamblott, et al., Derivation of Pluripotent Stem Cells from Cultured human Primordial Germ Cells, Proc. Natl. Acad. Sci. USA 95:13726 (1998)
	AY	Smith et al., Culture and Differentiation of Embryonic Stem Cells, J. Tiss. Cult. Meth. 13:89 (1991)
	AZ	Thomson, et al., Embryonic Stem Cell Lines Derived from Human Blastocysts, Science 282:1145 (1998)
	ВА	Trojanowski, et al., Transfectable and Transplantable Postmitotic Human Neurons: A Potential "Platform" for Gene Therapy of nervous System Diseases, Exp. Neurol. 144:92 (1997)
	ВВ	Tropepe, et al., Abstract 205.18: Autonomous Neural Cell Fate Specification in Mouse Embryonic Stem Cells, Soc. Neurosci. 25:527 (1999)
	вс	Tropepe, et al., Abstract 205.17: Neural Determination Genes Revealed by Expression Trapping in Embryonic Stem Cells, Soc. Neurosci. 25: 527 (1999)
	BD	Van Inzen, et al., Neuronal Differentiation of Embryonic Stem Cells, Biochim. Biophys. Acta 1312:21 (1996)
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	BF	Wojcik, et al., Catecholaminergic Neurons Result from Intracerebral Implantation of Embryonal Carcinoma Cells, Proc. Natl. Acad. Sci. USA 90:1305-130
	BG	Yandava, et al., "Global" Cell Replacement is Feasible Via Neural Stem Cell Transplantation: Evidence from the Dysmyelinated Shiverer Mouse Brain, Proc. Natl. Acad. Sci. USA 96:7029 (1999)
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